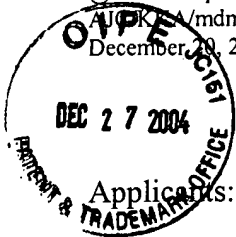


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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants: Garner T. Hauptert, Jr. and Steven R. Gullans  
Application No.: 10/601,184 Group Art Unit: 1644  
Filed: June 20, 2003 Examiner: Not yet assigned  
Confirmation No.: 6779  
Title: BIOSYNTHESIS OF HIF

CERTIFICATE OF MAILING OR TRANSMISSION	
I hereby certify that this correspondence is being deposited with the United States Postal Service with sufficient postage as First Class Mail in an envelope addressed to Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450, or is being facsimile transmitted to the United States Patent and Trademark Office on:	
December 20, 2004	
Date	Signature
Meredith Murray	
Typed or printed name of person signing certificate	

INFORMATION DISCLOSURE STATEMENT

Mail Stop Amendment  
Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

Sir:

This Information Disclosure Statement is submitted:

- ☐ under 37 CFR 1.129(a), or  
(First/Second submission after Final Rejection)
- ☒ under 37 CFR 1.97(b), or  
(Within any one of the following time periods: three months of filing national application (other than a CPA) or date of entry of the national stage in an international application; or before the mailing date of a first office action on the merits in a non-provisional application, including a CPA, or a Request for Continued Examination).
- ☐ under 37 CFR 1.97(c) together with either:
- ☐ a Statement under 37 CFR 1.97(e), as checked below, or
- ☐ a \$180.00 fee under 37 CFR 1.17(p), or  
(After the 37 CFR 1.97(b) time period, but before final action or notice of allowance, whichever occurs first)
- ☐ under 37 CFR 1.97(d) together with:
- ☐ a Statement under 37 CFR 1.97(e), as checked below, and
- ☐ a \$180.00 fee under 37 CFR 1.17(p), or  
(Filed after final action or notice of allowance, whichever occurs first, but on or before payment of the issue fee)
- ☐ under 37 CFR 1.97(i):  
Applicant requests that the IDS and cited reference(s) be placed in the application filewrapper.  
(Filed after payment of issue fee)

Statement Under 37 CFR 1.97(e)

- ☐ Each item of information contained in this Information Disclosure Statement was first cited in any communication from a foreign patent office in a counterpart foreign application not more than three months prior to the filing of this Information Disclosure Statement; or
- ☐ No item of information contained in this Information Disclosure Statement was cited in a communication from a foreign patent office in a counterpart foreign application, and, to the knowledge of the undersigned, after making reasonable inquiry, no item of information contained in the information disclosure statement was known to any individual designated in 37 CFR 1.56(c) more than three months prior to the filing of this Information Disclosure Statement.

Statement Under 37 CFR 1.704(d) (Patent Term Adjustment)

Applies to original applications (other than design) filed on or after May 29, 2000

- ☐ Each item of information contained in the Information Disclosure Statement was cited in a communication from a foreign patent office in a counterpart application and this communication was not received by any individual designated in § 1.56(c) more than thirty days prior to the filing of the Information Disclosure Statement.
- ☒ Enclosed herewith is form PTO-1449:
  - ☒ Copies of the cited references are enclosed (AA-AH, AL-AQ, AL2-AN2, AR-AZ, AR2-AZ2, AR3-AZ3, AR4-AV4).
    - ☐ Since this application was filed after June 30, 2003, copies of issued U.S. patents and published U.S. applications are not required and are not being provided.
  - ☐ Copies of the cited references are enclosed except those entered in prior application, U.S. Application No. [ ], to which priority under 35 U.S.C. 120 is claimed. [The earlier application contains copies of the cited references.]
  - ☒ The listed references (AA, AB, AP and AR4) were cited in the enclosed International Search Report in a counterpart foreign application.
  - ☐ The "concise explanation" requirement (non-English references) for reference(s) [ ] under 37 CFR 1.98(a)(3) is satisfied by:
    - ☐ the explanation provided on the attached sheet.
    - ☐ the explanation provided in the Specification.
    - ☐ submission of the enclosed International Search Report.
    - ☐ submission of the enclosed English-language version of a foreign Search Report and/or foreign Office Action.
    - ☐ the enclosed English language abstract.

[ X ] Applicant requests that the following non-published pending applications be considered:

Examiner's  
Initials

\_\_\_\_ U.S. Patent Application No. 09/220,913, by Garner T. Hauptert, Jr. ,  
filed December 24, 1998, Atty. Docket No.: 0838.1002-000

\_\_\_\_ U.S. Patent Application No. 09/412,268, by Behnaz Parharmi-Seren, *et al.*,  
filed October 5, 1999, Atty. Docket No.: 0838.1004-000

\_\_\_\_ U.S. Patent Application No. 09/506,246, by Rex T. Gallagher, *et al.*,  
filed February 17, 2000, Atty. Docket No.: 2075.1002-003

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Examiner

\_\_\_\_\_  
Date

- [ X ] A copy of each above-cited application, including the current claims, is enclosed.
- [ ] A copy of each above-cited application, including the current claims, is enclosed, except those entered in prior application, U.S. Application No. [ ], to which priority under 35 U.S.C. 120 is claimed.

The Examiner is requested to return a copy of the above list of pending applications indicating which references were considered with the next office communication.

It is requested that the information disclosed herein be made of record in this application.

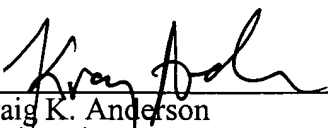
Method of payment:

- [ ] A check for the fee noted above is enclosed, or the fee has been included in the check with the accompanying Reply. A copy of this Statement is enclosed.
- [ ] Please charge Deposit Account 08-0380 in the amount of \$[ ]. A copy of this Statement is enclosed.
- [X] Please charge any deficiency in fees and credit any overpayment to Deposit Account 08-0380.

Respectfully submitted,

HAMILTON, BROOK, SMITH & REYNOLDS, P.C.

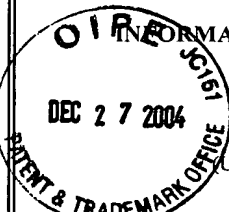
By

  
Kraig K. Anderson  
Registration No.: 54,961  
Telephone: (978) 341-0036  
Facsimile: (978) 341-0136

Concord, MA 01742-9133

Dated:

*December 20, 2004*

PTO-1449 REPRODUCED		ATTORNEY DOCKET NO. 0838.2003-001	APPLICATION NO. 10/601,184	
 <p>INFORMATION DISCLOSURE CITATION IN AN APPLICATION</p> <p>December 20, 2004</p> <p>(Use several sheets if necessary)</p>		APPLICANT Garner T. Hauptert, Jr., <i>et al.</i>		
		FILING DATE June 20, 2003	CONFIRMATION NO. 6779	GROUP 1644

## U.S. PATENT DOCUMENTS

EXAM- INER INI- TIAL	REF. NO.	DOCUMENT NUMBER	ISSUE DATE / PUBLICATION DATE	NAME
	AA	6,265,383 B1	July 24, 2001	Hauptert, Jr.
	AB	6,274,558 B1	August 14, 2001	Hauptert, Jr.
	AC	5,716,937	February 10, 1998	Hauptert, Jr.
	AD	5,910,484	June 8, 1999	Hauptert, Jr.
	AE	US 2004 - 0248080 A1	December 9, 2004	Hauptert, Jr.
	AF	5,844,091	December 1, 1998	Blaustein, <i>et al.</i>
	AG	5,695,756	December 9, 1997	Blaustein, <i>et al.</i>
	AH	5,164,296	November 17, 1992	Blaustein, <i>et al.</i>
	AI			
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EXAMINER	DATE CONSIDERED
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	APPLICANT Garner T. Hauptert, Jr., <i>et al.</i>			
	FILING DATE June 20, 2003		CONFIRMATION NO. 6779	GROUP 1644

FOREIGN PATENT DOCUMENTS						
		DOCUMENT NUMBER Country Code-Number-Kind Code (if known)	DATE MM-DD-YYYY	NAME OF PATENTEE OR APPLICANT OF CITED DOCUMENT	TRANSLATION YES NO	
	AL	WO 98/53832 A1	12/03/1998	The General Hospital Corporation		
	AM	WO 98/54195 A1	12/03/1998	The General Hospital Corporation, <i>et al.</i>		
	AN	WO 98/07739 A1	02/26/1998	The General Hospital Corporation, <i>et al.</i>		
	AO	WO 98/53831 A1	12/03/1998	The General Hospital Corporation		
	AP	WO 00/39328 A1	07/06/2000	The General Hospital Corporation		
	AQ	WO 01/25281 A1	04/12/2001	The General Hospital Corporation		
	AL2	WO 2004/000328 A1	12/31/2003	The General Hospital Corporation, <i>et al.</i>		
	AM2	EP 0 983 071 B1	03/08/2000	The General Hospital Corporation		
	AN2	EP 0 433 074 B1	06/19/1991	The General Hospital Corporation		
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OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)		
AR	Bennett, R.D., <i>et al.</i> , “ Progesterone Metabolism in <i>Digitalis Lanata</i> ”, <i>Phytochemistry</i> 7: 41-50 (1968).	
AS	Blaustein, M. P., “Sodium Ions, Calcium Ions, Blood Pressure Regulation, and Hypertension: A Reassessment and a Hypothesis”, <i>Am. J. Physiol.</i> , 232(5): C165-C173 (1977).	
AT	Blaustein, M.P., “Endogenous Ouabain: Role in the Pathogenesis of Hypertension”, <i>Kidney Int.</i> 49(6): 1748-1753 (1996).	
AU	Cantiello, <i>et al.</i> , “Na <sup>+</sup> Pump in Renal Tubular Cells is Regulated by Endogenous Na <sup>+</sup> , K <sup>+</sup> - ATPase Inhibitor from Hypothalamus”, <i>Am. J. Physiol.</i> , 255: F574-580 (1988).	
AV	Capsi, E. and Lewis, D.O., “Progesterone: Its Possible Role in the Biosynthesis of Cardenolides in <i>Digitalis Lanata</i> ”, <i>Science</i> 156(3774): 519-520 (1967).	
AW	Carilli, C. T., <i>et al.</i> , “Hypothalamic Factor Inhibits the (Na,K) ATPase from the Extracellular Surface”, <i>The Journal of Biological Chemistry</i> , 260(2): 1027-1031 (1985).	
AX	Crabos, M, <i>et al.</i> , “Atrial Natriuretic Peptide Regulates Release of Na <sup>+</sup> -K <sup>+</sup> -ATPase Inhibitor from Rat Brain”, <i>Am. J. Physiol.</i> , 254(6): F912-F917 (1988).	
AY	DeAngelis, C. and Hauptert, Jr., G. T., “Hypoxia Triggers Release of an Endogenous Inhibitor of Na <sup>+</sup> , -K <sup>+</sup> -ATPase from Midbrain and Adrenal”, <i>Am. J. Physiol.</i> , 274(1): F182-F188 (1998).	
AZ	Ferrandi, M., <i>et al.</i> , “Age-Dependency and Dietary Influence on the Hypothalamic Ouabain-Like Factor in Milan Hypertensive Rats”, <i>Journal of Hypertension</i> , 13(12): 1571-1574 (1995).	
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AS2	Ferrandi, M. and Manuta, P., “Ouabain-Like Factor: Is it the Natriuretic Hormone?”, <i>Curr. Opin. Nephrol. Hypertens.</i> 9(2): 165-171 (2000).	
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AU2	Goto, A., <i>et al.</i> , “Physiology and Pharmacology of Endogenous Digitalis-Like Factors”, <i>Pharmacological Reviews</i> , 44(3): 377-399 (1992).	

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OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)		
AV2	Haber, E. and Hauptert, Jr., G. T., "The Search for a Hypothalamic Na <sup>+</sup> , K <sup>+</sup> -ATPase Inhibitor", <i>Hypertension</i> , 9(4): 315-324 (1987).	
AW2	Haddy, F. J. and Overbeck, H. W., "The Role of Humoral Agents in Volume Expanded Hypertension", <i>Life Sciences</i> , 19(7): 935-948 (1976).	
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AY2	Hauptert, Jr., G. and Sancho, J. M., "Sodium Transport Inhibitor from Bovine Hypothalamus", <i>Proc. Natl. Acad. Sci.</i> , 76(9): 4658-4660 (1979).	
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AS3	Kawamura, A., <i>et al.</i> , "On the Structure of Endogenous Ouabain", <i>Proc. Natl. Acad. Sci.</i> , 96: 6654-6659 (1999).	
AT3	Kitano, S., <i>et al.</i> , "Exogenous Ouabain is Accumulated in the Adrenals and Mimics the Kinetics of Endogenous Digitalis-Like Factor in Rats", <i>Hypertens Res</i> , 21(1) : 47-56 (1998).	
AU3	Komiyama, Y., <i>et al.</i> , "Identification of Endogenous Ouabain in Culture Supernatant of PC12 Cells", <i>Journal of Hypertension</i> , 19(2): 229-236 (2001).	
AV3	Laredo, J., <i>et al.</i> , "Angiotensin II Stimulates Secretion of Endogenous Ouabain from Bovine Adrenocortical Cells via Angiotensin Type 2 Receptors", <i>Hypertension</i> , 29(1): 401-407 (1997).	
AW3	Lichtstein, D., <i>et al.</i> , "Biosynthesis of Digitalis-Like Compounds in Rat Adrenal Cells: Hydroxycholesterol as Possible Precursor", <i>Life Sciences</i> , 62(23): 2109-2126 (1998).	
AX3	Lu, Z., <i>et al.</i> , "Biosynthesis of Endogenous Ouabain from Pregnenolone and Progesterone in Bovine Adrenal Cortical Cells", <i>Hypertension</i> , 32(3): 624 (1998).	

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OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)		
AY3	Orth, D. N. and Kovacs, W. J., "The Adrenal Cortex". In <i>Williams Textbook of Endocrinology</i> 9th Edition, Jean. D. Wilson, M.D., <i>et al.</i> , eds. (W. B. Saunders Company), pp. 517-664 (1998).	
AZ3	Perrin, A., <i>et al.</i> , "Bovine Adrenocortical Cells in Culture Synthesize an Ouabain-Like Compound", <i>Molecular and Cellular Endocrinology</i> , 126: 7-15 (1997).	
AR4	Quazzaz, H. M. A. M., <i>et al.</i> , "Deglycosylated Products of Endogenous Digoxin-Like Immunoreactive Factor in Mammalian Tissue", <i>The Journal of Biological Chemistry</i> , 271(15): 8731-8737 (1996).	
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AV4	Elbashir, S.M., <i>et al.</i> , "Duplexes of 21-Nucleotide RNAs Mediate RNA Interference in Cultured Mammalian Cells," <i>Nature</i> , 411: 494-498 (2001).	

EXAMINER	DATE CONSIDERED
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